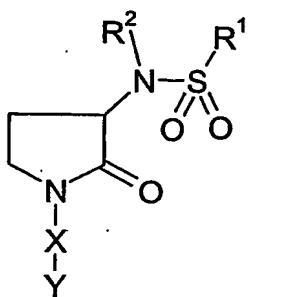


Claims

1. A compound of formula (I):

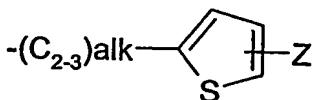
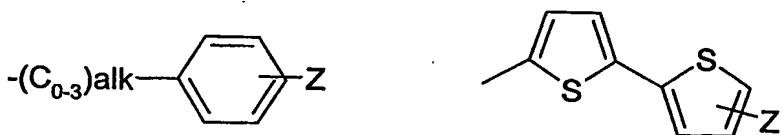
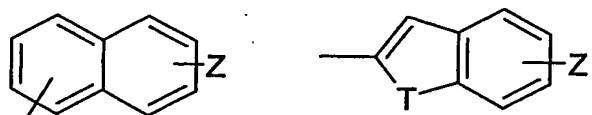


5

(I)

wherein:

R¹ represents a group selected from:



- each ring of which optionally contains a further heteroatom N,
10 Z represents an optional substituent halogen,
alk represents alkylene or alkenylene,
T represents S, O or NH;

R² represents hydrogen, -C₁₋₆alkyl, -C₁₋₃alkylCONR^aR^b, -C₁₋₃alkylCO₂C₁₋₄alkyl, -CO₂C₁₋₄alkyl or -C₁₋₃alkylCO₂H;

R^a and R^b independently represent hydrogen, -C₁₋₆alkyl, or together with the N atom to which they are bonded form a 5-, 6- or 7- membered non-aromatic heterocyclic ring

optionally containing an additional heteroatom selected from O, N or S, optionally substituted by C₁₋₄alkyl, and optionally the S heteroatom is substituted by O, i.e. represents S(O)_n;

5 n represents 0-2;

X represents phenyl or a 5- or 6- membered aromatic heterocyclic group containing at least one heteroatom selected from O, N or S, each of which is optionally substituted by 0-2 groups selected from: halogen, -C₁₋₄alkyl, -C₂₋₄alkenyl, -CN, -CF₃, -NR^aR^b, -C₀₋₄alkylOR^e, 10 -C(O)R^f and -C(O)NR^aR^b;

R^e represents hydrogen or -C₁₋₆alkyl;

R^f represents -C₁₋₆alkyl;

15

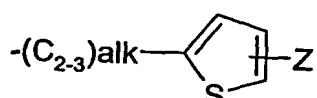
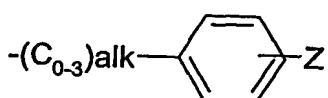
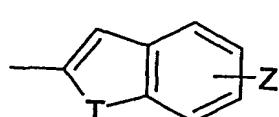
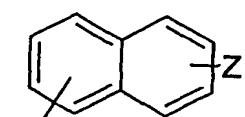
Y represents a group -C(R^x)(R^z)C₀₋₂alkylNR^cR^d,

R^x represents C₁₋₄alkyl optionally substituted by halogen;

20 R^z represents hydrogen or C₁₋₄alkyl optionally substituted by halogen;

R^c and R^d independently represent hydrogen, -C₁₋₆alkyl, -C₁₋₄alkylOH, or together with the N atom to which they are bonded form a 4-, 5-, 6- or 7- membered non-aromatic heterocyclic ring, the 5-, 6- or 7- membered non-aromatic heterocyclic ring optionally 25 containing an additional heteroatom selected from O, N or S, optionally substituted by C₁₋₄alkyl; and/or pharmaceutically acceptable derivative thereof.

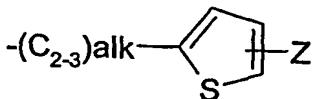
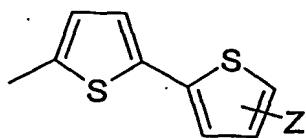
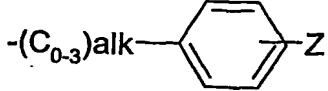
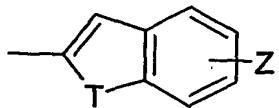
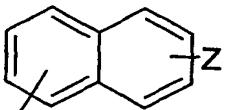
2. A compound according to claim 1 wherein R¹ represents a group selected from:



each ring of which optionally contains a further heteroatom N,
Z represents an optional substituent halogen,

alk represents alkylene or alkenylene,
 T represents S, O or NH
 and/or pharmaceutically acceptable derivative thereof.

- 5 3. A compound according to claim 1 or claim 2 wherein R² represents hydrogen and/or pharmaceutically acceptable derivative thereof.
4. A compound according to any one of claims 1-3 wherein X represents phenyl or a 5- or 6-membered aromatic heterocyclic group containing at least one heteroatom selected from O, N or S, each of which is optionally substituted by 0-2 groups selected from: halogen, -C₁₋₄alkyl and -NR^aR^b and/or pharmaceutically acceptable derivative thereof.
- 10 5. A compound according to any one of claims 1-4 wherein Y represents a group -C(R^x)(R^z)NR^cR^d and/or pharmaceutically acceptable derivative thereof.
- 15 6. A compound according to claim 1 wherein R¹ represents a group selected from:



each ring of which optionally contains a further heteroatom N,
 Z represents an optional substituent halogen,
 alk represents alkylene or alkenylene,

- 20 T represents S, O or NH;

R² represents hydrogen, -C₁₋₆alkyl, -C₁₋₃alkyCONR^aR^b, -C₁₋₃alkyICO₂C₁₋₄alkyl, -CO₂C₁₋₄alkyl or -C₁₋₃alkyICO₂H;

- 25 R^a and R^b independently represent hydrogen, -C₁₋₆alkyl, or together with the N atom to which they are bonded form a 5-, 6- or 7-membered non-aromatic heterocyclic ring

optionally containing an additional heteroatom selected from O, N or S, optionally substituted by C₁₋₄alkyl, and optionally the S heteroatom is substituted by O, i.e. represents S(O)_n;

5 X represents phenyl or a 5- or 6- membered aromatic heterocyclic group containing at least one heteroatom selected from O, N or S, each of which is optionally substituted by 0-2 groups selected from: halogen, -C₁₋₄alkyl, -C₂₋₄alkenyl, -CN, -CF₃, -NR^aR^b, -C₀₋₄alkylOR^e, -C(O)R^f and -C(O)NR^aR^b;

10 R^e represents hydrogen or -C₁₋₆alkyl;

R^f represents -C₁₋₆alkyl;

Y represents a group -C(R^x)(R^z)C₀₋₂alkylNR^cR^d;

15

R^x represents C₁₋₄alkyl optionally substituted by halogen (e.g. CF₃, -CH₂CF₃);

R^z represents hydrogen or C₁₋₄alkyl optionally substituted by halogen (e.g. CF₃, -CH₂CF₃);

20 R^c and R^d independently represent hydrogen, -C₁₋₆alkyl, -C₁₋₄alkylOH, or together with the N atom to which they are bonded form a 5- or 6- membered non-aromatic heterocyclic ring optionally containing an additional heteroatom selected from O, N or S, optionally substituted by C₁₋₄alkyl;
and pharmaceutically acceptable derivatives thereof.

25

7. A compound according to claim 1 selected from:

(E)-2-(5-Chloro-2-thienyl)-N-(1-{4-[1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;

(E)-2-(5-Chloro-2-thienyl)-N-(1-{2-fluoro-4-[1-(4-morpholinyl)ethyl]phenyl}-2-oxo-3-

30 pyrrolidinyl)ethenesulfonamide;

(E)-2-(5-Chloro-2-thienyl)-N-[1-(2-fluoro-4-{1-[(2-hydroxyethyl)(methyl)amino]ethyl}phenyl)-2-oxo-3-pyrrolidinyl]ethenesulfonamide;

(E)-N-{1-[4-(1-Aminoethyl)-2-fluorophenyl]-2-oxo-3-pyrrolidinyl}-2-(5-chloro-2-thienyl)ethenesulfonamide;

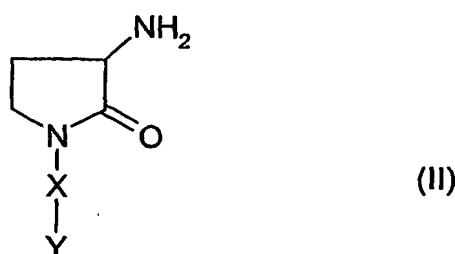
35 6-Chloro-N-(1-{4-[1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-1-benzothiophene-2-sulfonamide;

(E)-2-(5-Chloro-2-thienyl)-N-((3S)-1-{4-[(1S)-1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;

- 6-Chloro-N-((3*S*)-1-{4-[(1*S*)-1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-2-naphthalenesulfonamide;
- 6-Chloro-N-((3*S*)-1-{4-[(1*S*)-1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-1-benzothiophene-2-sulfonamide;
- 5 (*E*)-2-(5-Chloro-2-thienyl)-N-((3*S*)-1-{4-[(1*R*)-1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;
- 6-Chloro-N-((3*S*)-1-{4-[(1*R*)-1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-2-naphthalenesulfonamide;
- 6-Chloro-N-((3*S*)-1-{4-[(1*R*)-1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-1-benzothiophene-2-sulfonamide;
- (*E*)-2-(5-Chloro-2-thienyl)-N-((3*S*)-1-{4-[(1*R*)-1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;
- (*E*)-2-(5-Chloro-2-thienyl)-N-((3*S*)-1-{4-[(1*S*)-1-(dimethylamino)ethyl]-2,6-difluorophenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;
- 15 (*E*)-2-(5-Chloro-2-thienyl)-N-((3*S*)-1-{4-[1-(dimethylamino)propyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;
- (*E*)-2-(5-Chloro-2-thienyl)-N-((3*S*)-1-{4-[1-(dimethylamino)-2-methylpropyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;
- (*E*)-2-(5-Chloro-2-thienyl)-N-(1-{4-[1-(dimethylamino)-1-methylethyl]-2-fluorophenyl}-2-
- 20 oxo-3-pyrrolidinyl)ethenesulfonamide;
- 6-Chloro-N-(1-{4-[1-(dimethylamino)-1-methylethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-2-naphthalenesulfonamide;
- 6-Chloro-N-(1-{4-[1-(dimethylamino)-1-methylethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-1-benzothiophene-2-sulfonamide;
- 25 6-Chloro-N-(1-{4-[1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-2-naphthalenesulfonamide;
- 6-Chloro-N-(1-{4-[1-(ethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-2-naphthalenesulfonamide;
- 6-Chloro-N-[1-(4-{1-[ethyl(methyl)amino]ethyl}-2-fluorophenyl)-2-oxo-3-pyrrolidinyl]-2-
- 30 naphthalenesulfonamide;
- 6-Chloro-N-[1-(2-fluoro-4-{1-[(1-methylethyl)amino]ethyl}phenyl)-2-oxo-3-pyrrolidinyl]-2-naphthalenesulfonamide;
- 6-Chloro-N-[1-(2-fluoro-4-{1-[methyl(1-methylethyl)amino]ethyl}phenyl)-2-oxo-3-
- 35 pyrrolidinyl]-2-naphthalenesulfonamide;
- N-(1-{4-[1-(1-Azetidinyl)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-6-chloro-2-naphthalenesulfonamide;

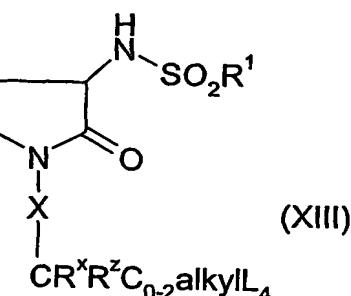
- 6-Chloro-N-(1-{2-fluoro-4-[1-(1-pyrrolidinyl)ethyl]phenyl}-2-oxo-3-pyrrolidinyl)-2-naphthalenesulfonamide;
- 6-Chloro-N-(1-{2-fluoro-4-[1-(1-piperidinyl)ethyl]phenyl}-2-oxo-3-pyrrolidinyl)-2-naphthalenesulfonamide;
- 5 5'-Chloro-N-((3S)-1-{4-[1-(dimethylamino)ethyl]-2-fluorophenyl}-2-oxo-3-pyrrolidinyl)-2,2'-bithiophene-5-sulfonamide;
- (E)-2-(5-Chloro-2-thienyl)-N-((3S)-1-{4-[(1S)-1-(dimethylamino)ethyl]phenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;
- (E)-2-(5-Chloro-2-thienyl)-N-((3S)-1-{4-[(1R)-1-(dimethylamino)ethyl]phenyl}-2-oxo-3-pyrrolidinyl)ethenesulfonamide;
- 10 6-Chloro-N-((3S)-1-{4-[(1S)-1-(dimethylamino)ethyl]phenyl}-2-oxo-3-pyrrolidinyl)-1-benzothiophene-2-sulfonamide;
- 6-Chloro-N-((3S)-1-{4-[(1R)-1-(dimethylamino)ethyl]phenyl}-2-oxo-3-pyrrolidinyl)-1-benzothiophene-2-sulfonamide;
- 15 (1E)-2-(5-Chloro-2-thienyl)-N-(1-{4-[1-(dimethylamino)ethyl]-2,6-difluorophenyl}-2-oxo-3-pyrrolidinyl)-1-propene-1-sulfonamide;
- 6-Chloro-N-(1-{4-[1-(dimethylamino)ethyl]-2,6-difluorophenyl}-2-oxo-3-pyrrolidinyl)-1-benzothiophene-2-sulfonamide;
- and/or pharmaceutically acceptable derivative thereof.
- 20 8. A compound according to any one of claims 1-7 and/or pharmaceutically acceptable derivative thereof for use in therapy.
9. A pharmaceutical composition comprising a compound according to any one of claims
- 25 1-7 and/or pharmaceutically acceptable derivative thereof together with at least one pharmaceutical carrier and/or excipient.
10. Use of a compound according to any one of claims 1-7 and/or pharmaceutically acceptable derivative thereof for the manufacture of a medicament for the treatment of a
- 30 patient suffering from a condition susceptible to amelioration by a Factor Xa inhibitor.
11. A method of treating a patient suffering from a condition susceptible to amelioration by a Factor Xa inhibitor comprising administering a therapeutically effective amount of a compound according to any one of claims 1-7 and/or pharmaceutically acceptable derivative thereof.
- 35 12. A process for preparing a compound of formula (I) which comprises:

(a) reacting compounds of formula (II) or an acid addition salt thereof with compounds of formula (III) where V is a suitable leaving group:



OR:

(b) by reacting compounds of formula (XIII) with HNR^cR^d:



10 OR:

(c) by reacting compounds of formula (I) where R² is hydrogen with compounds of formula (XVII):

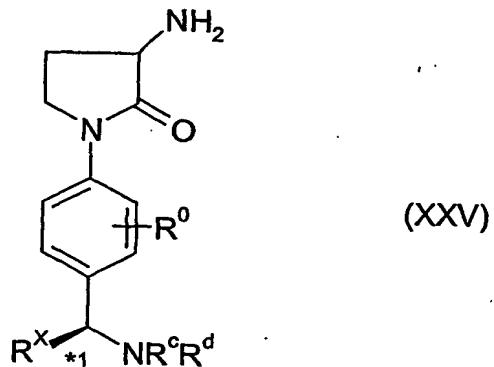


wherein R² is -C₁₋₆alkyl, -C₁₋₃alkylCONR^aR^b, -C₁₋₃alkylCO₂C₁₋₄alkyl, or -CO₂C₁₋₄alkyl and T is a suitable leaving group, optionally followed by removal of the alkyl protecting group where appropriate;

20

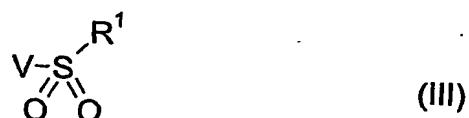
OR:

(d) by reacting a compound of formula (XXV) where X represents phenyl, Y represents $\text{CH}(\text{R}^x)\text{NR}^c\text{R}^d$, R^c and R^d each represent the same C_{1-6} alkyl substituent and R^0 represents 0-2 optional substituents on the phenyl ring selected from: halogen, $-\text{C}_{1-4}$ alkyl, $-\text{C}_{2-4}$ alkenyl, $-\text{CN}_1$, $-\text{CF}_3$, $-\text{NR}^a\text{R}^b$, $-\text{C}_{0-4}\text{alkylOR}^e$, $-\text{C}(\text{O})\text{R}^f$ and $\text{C}(\text{O})\text{NR}^a\text{R}^b$ and/or an acid addition salt 5 thereof:



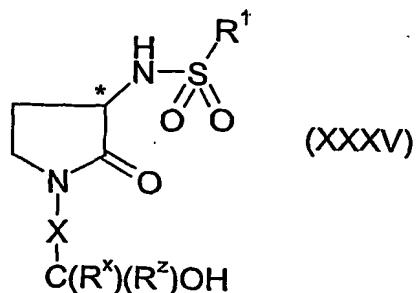
with a compound of formula (III) where V is a suitable leaving group:

10



OR:

(e) treatment of compounds of formula (XXXV) where Y represents $-\text{C}(\text{R}^x)(\text{R}^z)\text{NR}^c\text{R}^d$ and 15 R^x and R^z both represent C_{1-4} alkyl and R^2 represents hydrogen:

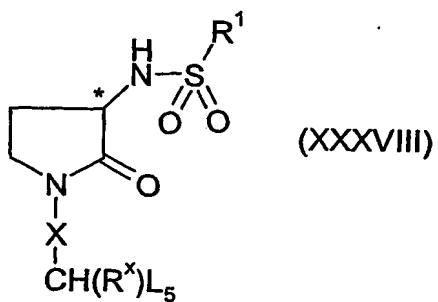


with hydrogen chloride in the presence of zinc chloride, followed by reaction with HNRR^cR^d ; 20

OR:

(f) by reacting compounds of formula (XXXVIII) where Y represents $-C(R^x)NR^cR^d$, R^x represents C_{1-4} alkyl and R^c and R^d independently represent hydrogen, C_{1-6} alkyl, or together with the N atom to which they are bonded form a 4-, 5-, 6- or 7- membered non-aromatic heterocyclic ring and L_5 is a suitable leaving group:

5



with $HNRR^cR^d$.